

## WEST Search History

DATE: Tuesday, April 06, 2004

<u>Hide?</u>	<u>Set Name</u>	<u>Query</u>	<u>Hit Count</u>
<i>DB=PGPB,USPT,EPAB,JPAB,DWPI; PLUR=YES; OP=ADJ</i>			
<input type="checkbox"/>	L8	l3 and l4	4
<input type="checkbox"/>	L7	l6 and rhodamin\$6.ab.	6
<input type="checkbox"/>	L6	l4 and therap\$6	139
<input type="checkbox"/>	L5	l3 and l4L4	0
<input type="checkbox"/>	L4	l2 near5 salt	485
<input type="checkbox"/>	L3	photodynamic and L2	329
<input type="checkbox"/>	L2	rhodamine\$	25309
<input type="checkbox"/>	L1	us-5556992-\$ did.	2

END OF SEARCH HISTORY

ROY 10/088,072

=> d his

(FILE 'HOME' ENTERED AT 14:53:13 ON 06 APR 2004)

FILE 'HCAPLUS' ENTERED AT 14:53:21 ON 06 APR 2004

L1       19 S GUIMOND M?/AU  
L2       5 S MOLFINO N?/AU  
L3       1784 S ROY D?/AU  
L4       1805 S L1-3  
L5       3 S L4 AND RHODAMINE  
          SELECT RN L5 2

FILE 'REGISTRY' ENTERED AT 14:54:53 ON 06 APR 2004

L6       6 S E1-6

FILE 'HCAPLUS' ENTERED AT 14:55:37 ON 06 APR 2004

L7       1 S L6 AND L5  
L8       13213 S L6  
L9       113 S L8 AND RHODAMINE  
L10      1 S L9 AND PHOTODYNAMIC  
L11      2 S L9 AND DIAGNOSTS  
L12      5474 S L8(L)THU/RL  
L13      26 S L9 AND L12  
L14      14 S L13 AND PY<2000

FILE 'REGISTRY' ENTERED AT 16:01:17 ON 06 APR 2004

L15     5 S L6 NOT 59865-13-3

FILE 'HCAPLUS' ENTERED AT 16:04:54 ON 06 APR 2004

L16     1 S L15  
L17     0 S L16 NOT L5

FILE 'REGISTRY' ENTERED AT 16:05:41 ON 06 APR 2004

L18     STR 333957-98-5  
L19     0 S L18  
L20     52360 S OCS-C6-C6/ES  
L21     0 S L18 SSS SAM SUB=L20  
L22     STR L18  
L23     42 S L22  
L24     794 S L22 FUL  
          SAVE L24 ROY072P/A  
L25     0 S L18 SSS SAM SUB=L24  
L26     STR L18  
L27     0 S L26 SSS SAM SUB=L24  
L28     12 S L26 SSS FUL SUB=L24  
          SAVE L28 ROY072S1/A

FILE 'HCAPLUS' ENTERED AT 16:11:51 ON 06 APR 2004

L29     24 S L28  
L30     3946 S PHOTODYNAMIC THERAPY+PFT/CT  
L31     5182 S "PHOTOSENSITIZERS (PHARMACEUTICAL)"+PFT/CT  
L32     162992 S LYMPHOCYTE+PFT,NT/CT  
L33     35145 S "TRANSPLANT AND TRANSPLANTATION"+PFT/CT  
L34     59562 S IMMUNITY+PFT,NT/CT  
L35     10647 S RHODAMIN?/OBI  
L36     23 S L35 AND L30  
L37     12 S L30(L)RHODAMIN?  
L38     9 S L37 NOT L29  
          SELECT RN L38 4

FILE 'REGISTRY' ENTERED AT 16:23:29 ON 06 APR 2004

L39     10 S E7-16

FILE 'HCAPLUS' ENTERED AT 16:24:10 ON 06 APR 2004

L40     6 S L38 AND L39

ROY 10/088,072

L41 1 S L40 AND PATENT/DT  
L42 45210 S L39

FILE 'REGISTRY' ENTERED AT 16:27:56 ON 06 APR 2004  
L43 0 S 333957-95-2/CRN  
L44 44 S L24 AND BR>1  
L45 39 S L44 AND N=2  
L46 32 S L45 NOT IN/ELS  
L47 24 S L46 AND BR<4  
L48 10 S L47 AND "4,5-DIBROMO"  
L49 0 S L44 AND L39  
L50 44 S L44 AND L24  
L51 75 S L24 AND BR/ELS  
L52 0 S L51 AND L39  
L53 5 S L39 AND BR/ELS  
L54 0 S "BENZOIC ACID, 2-(3,6-DIETHYLAMINO-4,5-DIBROMO-9H-XANTHEN-9-  
L55 307 S OC5-C6-C6/ES AND 46.150.18/RID AND N=2 AND BR/ELS  
L56 65 S L55 AND BR=2  
L57 46 S L56 AND NRS=2  
L58 2 S L57 AND O=1  
L59 31 S L57 AND O=3  
L60 17 S L59 AND "4,5-DIBROMO"  
L61 4 S "2-(3,6-DIAMINO-4,5-DIBROMO-9H-XANTHEN-9-YL)"  
L62 2 S "4,5-DIBROMO" AND "BIS(DIETHYLAMINO)"  
L63 5 S L39 AND BR=2

FILE 'HCAPLUS' ENTERED AT 16:41:44 ON 06 APR 2004  
L64 2 S L62  
L65 2 S L63  
L66 3 S L64-65  
S L24 AND BR/ELS

FILE 'REGISTRY' ENTERED AT 17:06:41 ON 06 APR 2004  
L67 1223588 S BR/ELS

FILE 'HCAPLUS' ENTERED AT 17:06:42 ON 06 APR 2004  
FILE 'REGISTRY' ENTERED AT 17:06:49 ON 06 APR 2004  
L68 75 S L24 AND BR/ELS  
L69 32 S L68 AND "DIETHYL"  
L70 26 S L69 AND N=2  
L71 57 S L55 AND "BIS(DIETHYLAMINO)"  
L72 57 S L71 AND N=2  
L73 2 S L72 AND "4,5-DIBROMO"  
L74 1 S L73 AND CL/ELS  
L75 STR 333957-96-3  
L76 0 S L75 FAM  
L77 1 S L75 FAM FUL

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(FILE 'HOME' ENTERED AT 14:53:13 ON 06 APR 2004)

FILE 'HCAPLUS' ENTERED AT 14:53:21 ON 06 APR 2004

L1           19 S GUIMOND M?/AU  
 L2           5 S MOLFINO N?/AU  
 L3           1784 S ROY D?/AU  
 L4           1805 S L1-3  
 L5           3 S L4 AND RHODAMINE  
              SELECT RN L5 2

FILE 'REGISTRY' ENTERED AT 14:54:53 ON 06 APR 2004

L6           6 S E1-6

FILE 'HCAPLUS' ENTERED AT 14:55:37 ON 06 APR 2004

L7           1 S L6 AND LS

=> d ibib abs hitstr ind

L7 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2001:265273 HCAPLUS  
 DOCUMENT NUMBER: 134:292146  
 TITLE: Rhodamine derivatives for photodynamic  
         diagnosis and treatment  
 INVENTOR(S): Roy, Denis-Claude; Guimond, Martin  
               ; Molfino, Nestor A.  
 PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital  
                   Maisonneuve-Rosemont  
 SOURCE: PCT Int. Appl., 60 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

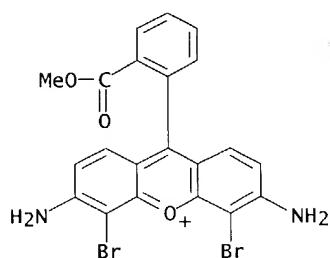
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
BR 2000014135	A	20020521	BR 2000-14135	20001003
EP 1267931	A1	20030102	EP 2000-965683	20001003
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
JP 2003510372	T2	20030318	JP 2001-527823	20001003
PRIORITY APPLN. INFO.:			US 1999-157790P P	19991005
			WO 2000-CA1142 W	20001003

AB The present invention relates to the use of the photoactivatable derivs. for  
 the photodynamic treatment for the selective destruction and/or  
 inactivation of immunol. reactive cells without affecting the normal cells  
 and without causing systemic toxicity for the patient, wherein appropriate  
 intracellular levels of said derivs. are achieved and irradn. of a  
 suitable wavelength and intensity is applied. Examples are given of the  
 selective phototoxicity of rhodamine derivs. against K562 cells,  
 CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and  
 human B cells. Immunol. disorders, including graft-vs-host disease are  
 treated with photodynamic therapy.

IT 333957-97-4  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological

process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (rhodamine derivs. for photodynamic diagnosis and treatment  
 of immunol. disorders)

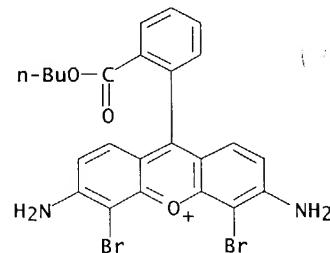
RN 333957-97-4 HCPLUS  
 CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-,  
 bromide (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

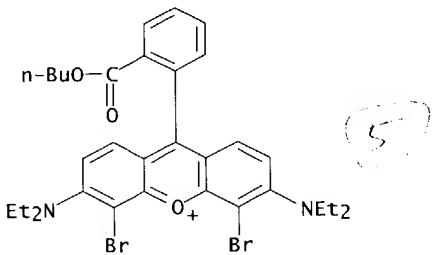
IT 333957-95-2 333957-96-3  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (rhodamine derivs. for photodynamic diagnosis and treatment  
 of immunol. disorders)

RN 333957-95-2 HCPLUS  
 CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, bromide  
 (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

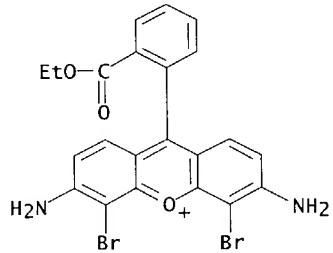
RN 333957-96-3 HCPLUS  
 CN Xanthylum, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-,  
 chloride (9CI) (CA INDEX NAME)

● Cl<sup>-</sup>

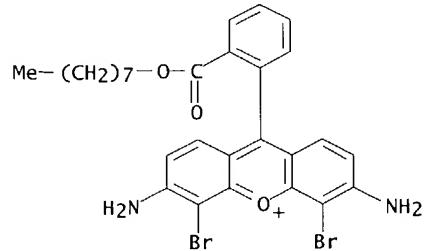
IT 333957-98-5 333957-99-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(rhodamine derivs. for photodynamic diagnosis and treatment  
of immunol. disorders)

RN 333957-98-5 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-, bromide  
(9CI) (CA INDEX NAME)● Br<sup>-</sup>

RN 333957-99-6 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-[(octyloxy)carbonyl]phenyl]-,  
bromide (9CI) (CA INDEX NAME)● Br<sup>-</sup>

IT 59865-13-3, Cyclosporin A

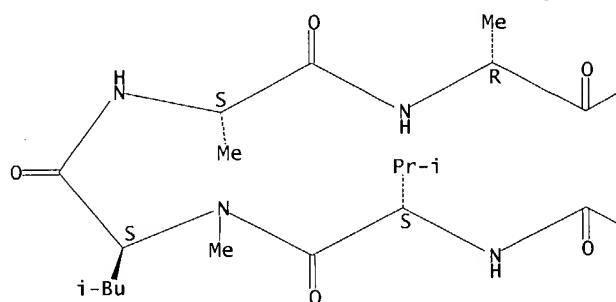
ROY 10/088,072

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
(rhodamine derivs. for photodynamic diagnosis and treatment  
of immunol. disorders: effect of cyclosporin A on rhodamine  
cellular efflux)

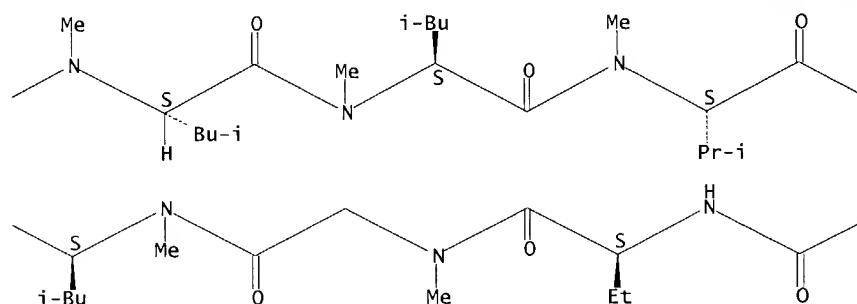
RN 59865-13-3 HCPLUS  
CN Cyclosporin A (9CI) (CA INDEX NAME)

Absolute stereochemistry.  
Double bond geometry as shown.

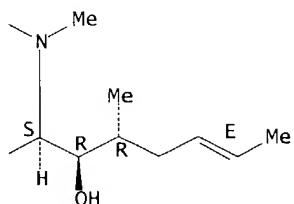
PAGE 1-A



PAGE 1-B



PAGE 1-C



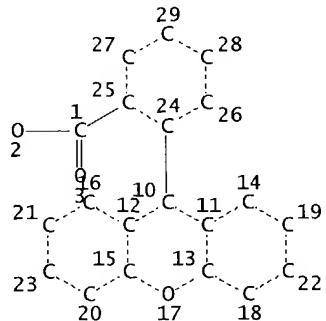
IC ICM A61K041-00  
CC 8-9 (Radiation Biochemistry)  
Section cross-reference(s): 15

ST **rhodamine deriv immunol disorder photodynamic therapy**  
 IT Lymphocyte  
     (PHA-activated; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: phototoxicity against immunol. reactive cells)  
 IT Immunity  
     (disorder; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT Cytometry  
     (flow; photoactivatable **rhodamine** derivs. for evaluating transport mechanism of cells by flow cytometry)  
 IT Transplant and Transplantation  
     (graft-vs.-host reaction; **rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT Allergy  
     Autoimmune disease  
     Diagnosis  
     Photodynamic therapy  
     Photosensitizers (pharmaceutical)  
     Transplant rejection  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT B cell (Lymphocyte)  
     CD4-positive T cell  
     CD8-positive T cell  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: phototoxicity against immunol. reactive cells)  
 IT Bone marrow  
     Hematopoietic precursor cell  
     Mononuclear cell (Leukocyte)  
     Transplant and Transplantation  
         (**rhodamine** derivs. for photodynamic ex vivo treatment of hematopoietic cells)  
 IT 333957-97-4  
     RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT 333957-95-2 333957-96-3  
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT 333957-98-5 333957-99-6  
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders)  
 IT 59865-13-3, Cyclosporin A  
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)  
         (**rhodamine** derivs. for photodynamic diagnosis and treatment of immunol. disorders: effect of cyclosporin A on **rhodamine** cellular efflux)

REFERENCE COUNT:       8       THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ROY 10/088,072

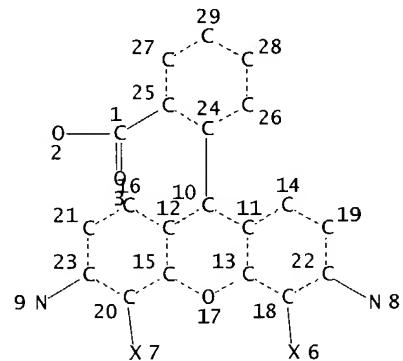
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L22 STR



NODE ATTRIBUTES:  
DEFAULT MLEVEL IS ATOM  
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
RSPEC I  
NUMBER OF NODES IS 23

STEREO ATTRIBUTES: NONE  
L24 794 SEA FILE=REGISTRY SSS FUL L22  
L26 STR



NODE ATTRIBUTES:  
CONNECT IS X4 RC AT 8  
CONNECT IS X4 RC AT 9  
DEFAULT MLEVEL IS ATOM  
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:  
RSPEC I  
NUMBER OF NODES IS 27

STEREO ATTRIBUTES: NONE  
L28 12 SEA FILE=REGISTRY SUB=L24 SSS FUL L26  
L29 24 SEA FILE=HCAPLUS ABB=ON PLU=ON L28

=> d ibib abs hitstr 1-24

L29 ANSWER 1 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN  
ACCESSION NUMBER: 2003:749309 HCAPLUS  
DOCUMENT NUMBER: 140:213073  
TITLE: Ca2+-dependent and caspase-3-independent apoptosis

caused by damage in Golgi apparatus due to  
2,4,5,7-tetrabromorhodamine 123 bromide-induced  
photodynamic effects

AUTHOR(S): Ogata, Maiko; Inanami, Osamu; Nakajima, Mihoko;  
Nakajima, Takayuki; Hiraoka, Wakako; Kuwabara,  
Mikinori

CORPORATE SOURCE: Laboratory of Radiation Biology, Department of  
Environmental Veterinary Science, Graduate School of  
Veterinary Medicine, Hokkaido University, Sapporo,  
Japan

SOURCE: Photochemistry and Photobiology (2003), 78(3), 241-247  
CODEN: PHCBAP; ISSN: 0031-8655

PUBLISHER: American Society for Photobiology

DOCUMENT TYPE: Journal

LANGUAGE: English

AB To clarify the role of the Golgi app. in photodynamic therapy-induced apoptosis, its signaling pathway was studied after photodynamic treatment of human cervix carcinoma cell line HeLa, in which a photosensitizer, 2,4,5,7-tetrabromorhodamine 123 bromide (TBR), was incorporated into the Golgi app. Laser scanning microscopic anal. of TBR-loaded HeLa cells confirmed that TBR was exclusively located in the Golgi app. HeLa cells incubated with TBR for 1 h were then exposed to visible light using an Xe lamp. Light of wavelength below 670 nm was eliminated with a filter. Morphol. observation of nuclei stained with Hoechst 33342 revealed that apoptosis of cells was induced by exposure to light. ESR spectrometry showed that light-exposed TBR produced both singlet oxygen (1O<sub>2</sub>) and superoxide anion (O<sub>2</sub><sup>-</sup>). Apoptosis induction by TBR was inhibited by pyrrolidine dithiocarbamate, an O<sub>2</sub><sup>-</sup> scavenger, but not by Na<sub>3</sub>, a quencher of 1O<sub>2</sub>. Furthermore, TBR-induced apoptosis was inhibited by aurintricarboxylic acid and ZnCl<sub>2</sub>, which are known as inhibitors of DNase (DNase) .gamma., and (acetoxymethyl)-1,2-bis(o-aminophenoxy)-ethane-N,N,N',N'-tetraacetic acid, a chelator of Ca<sup>2+</sup>, but not by acetyl Asp-Glu-Val-Asp-aldehyde, an inhibitor of caspase-3. These results suggested that O<sub>2</sub><sup>-</sup> was responsible for TBR-induced apoptosis, and Ca<sup>2+</sup>-dependent and caspase-3-independent nuclease such as DNase .gamma. played an important role in apoptotic signaling triggered by Golgi dysfunction.

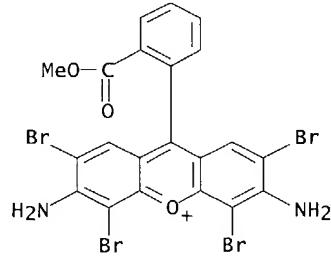
IT 623903-26-4, Tetrabromorhodamine 123

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tetrabromorhodamine 123; Golgi app., calcium, caspase, and DNase role in PDT-induced apoptosis in cervical carcinoma)

RN 623903-26-4 HCPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 2 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2003:226526 HCPLUS  
 DOCUMENT NUMBER: 139:377312  
 TITLE: "Mitochondrial" photochemical drugs do not release toxic amounts of 102 within the mitochondrial matrix space  
 AUTHOR(S): Petrat, Frank; Pindiur, Stanislaw; Kirsch, Michael; de Groot, Herbert  
 CORPORATE SOURCE: Institut fur Physiologische Chemie, Universitaetsklinikum, Essen, D-45122, Germany  
 SOURCE: Archives of Biochemistry and Biophysics (2003), 412(2), 207-215  
 CODEN: ABBIA4; ISSN: 0003-9861  
 PUBLISHER: Elsevier Science  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

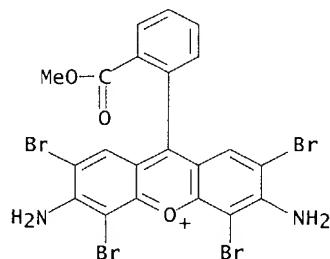
AB Previously, we demonstrated that mitochondrial NAD(P)H is the primary target of singlet oxygen (102) generated by photoactivation of mitochondria-selective rhodamine derivs. Hence, local NAD(P)H oxidn./fluorescence decrease may be used to reveal the site of intracellular 102 generation. Therefore, in addn. to the previously used tetramethylrhodamine methylester (TMRM), 2',4',5',7'-tetrabromorhodamine 123 bromide (TBRB) and rhodamine 123 (Rho 123), we tested here whether mitochondrial NAD(P)H of cultured hepatocytes is directly oxidized upon irradn. of different "mitochondrial" photosensitizers (Photofrin; protoporphyrin IX; Al(III) phthalocyanine chloride tetrasulfonic acid; meso-tetra(4-sulfonatophenyl)porphine dihydrochloride; Visudyne). In contrast to TMRM and Rho 123, which directly oxidized NAD(P)H upon irradn., irradn. of intracellular TBRB and the photochem. drugs only indirectly affected mitochondrial NAD(P)H due to loss of mitochondrial integrity. In line with this result only TMRM and Rho 123 exclusively localized within the mitochondrial matrix. Due to these results it is doubtful whether real mitochondrial photosensitizers actually exist among the photochem. drugs applicable/used for photodynamic therapy.

IT 623903-26-4

RL: PAC (Pharmacological activity); BIOL (Biological study)  
 (mitochondrial PDT photosensitizers do not release toxic amts. of 102 within mitochondrial matrix space)

RN 623903-26-4 HCPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

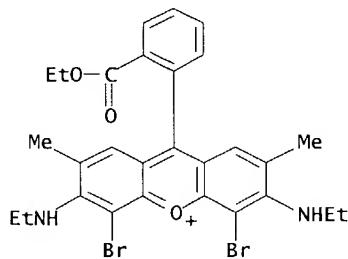
REFERENCE COUNT: 56 THERE ARE 56 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 3 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2002:777919 HCPLUS  
 DOCUMENT NUMBER: 137:280622

TITLE: Halogenated rhodamine dye derivatives and their therapeutic applications  
 INVENTOR(S): Habi, Abdelkrim; Gravel, Denis; Villeneuve, Luc;  
 Forte, Jean-Pierre; Su, Hongsheng; Vaillancourt, Marc  
 PATENT ASSIGNEE(S): Theratechnologies Inc., Can.  
 SOURCE: PCT Int. Appl., 117 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002079183	A1	20021010	WO 2002-CA438	20020327
WO 2002079183	C1	20030220		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
EP 1276734	A1	20030122	EP 2002-708105	20020327
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
BR 2002004489	A	20030401	BR 2002-4489	20020327
US 2003212126	A1	20031113	US 2003-297088	20030530
PRIORITY APPLN. INFO.: CA 2001-2342675 A 20010402 US 2001-822223 A 20010402 WO 2002-CA438 W 20020327				

OTHER SOURCE(S): MARPAT 137:280622  
 AB Bromo derivs. of rhodamine 110, rhodamine B, and rhodamine 6G and other halo rhodamine derivs. are useful as intermediates and as bactericides and antiviral agents and in the treatment of immunol. disorders. In an example, rhodamine B Me ester was dihydrogenated and then brominated and oxidized and treated with acetic acid to provide the purple acetate salt of 2,7-dibromorhodamine B Me ester.  
 IT 467232-05-9P  
 RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  
 (green-red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)  
 RN 467232-05-9 HCPLUS  
 CN Xanthylium, 4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-3,6-bis(ethylamino)-2,7-dimethyl-, bromide (9CI) (CA INDEX NAME)

● Br<sup>-</sup>

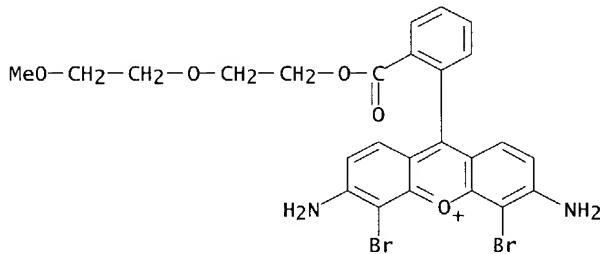
IT 467232-07-1P 467232-23-1P

RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)

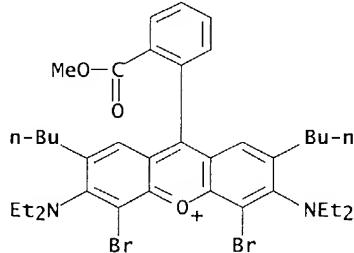
RN 467232-07-1 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-[[2-(2-methoxyethoxy)ethoxy]carbonyl]phenyl]-, bromide (9CI) (CA INDEX NAME)

● Br<sup>-</sup>

RN 467232-23-1 HCPLUS

CN Xanthylium, 4,5-dibromo-2,7-dibutyl-3,6-bis(diethylamino)-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)

● Br<sup>-</sup>

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

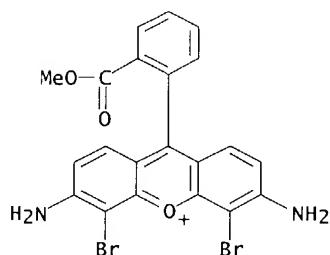
L29 ANSWER 4 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2002:541337 HCPLUS  
 DOCUMENT NUMBER: 137:167817  
 TITLE: P-glycoprotein targeting: a unique strategy to selectively eliminate immunoreactive T cells  
 AUTHOR(S): Guimond, Martin; Balassy, Antonia; Barrette, Melanie; Brochu, Sylvie; Perreault, Claude; Roy, Denis Claude  
 CORPORATE SOURCE: Division of Hematology-Immunology, Maisonneuve-Rosemont Hospital Research Center, Department of Medicine, Universite de Montreal, Montreal, QC, Can.  
 SOURCE: Blood (2002), 100(2), 375-382  
 CODEN: BLOOAW; ISSN: 0006-4971  
 PUBLISHER: American Society of Hematology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB T lymphocytes have been found to harbor P-glycoprotein (Pgp) and to demonstrate modulation of its ion channel transporter function according to the state of activation of T lymphocytes. We hypothesized that cytotoxic chems. that are extruded by Pgp could be used to specifically eliminate immunoreactive T-cell populations. In this study, we evaluated the capacity of 4,5-dibromorhodamine Me ester (TH9402), a photosensitizer structurally similar to rhodamine, a dye transported by Pgp, and which becomes highly cytotoxic on activation with visible light to selectively deplete alloreactive T lymphocytes. Stimulation of T cells with mitogens or allogeneic major histocompatibility complex-mismatched cells resulted in the preferential retention of the TH9402 rhodamine-deriv. in activated T cells, both CD4+ and CD8+. Photodynamic cell therapy of TH9402-exposed T cells led to the selective elimination of immunoreactive T-cell populations. In addn., this treatment preserved resting T cells and their capacity to respond to third-party cells. Inhibition of Pgp enhanced cellular trapping of the dye in nonactivated T cells and resulted in their depletion after exposure to light. Targeting of Pgp-deficient cells may therefore represent an appealing strategy for the prevention and treatment of graft-vs.-host disease and other alloimmune or autoimmune disorders.

IT 174230-05-8, TH9402  
 RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);  
 THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (P-glycoprotein targeting to selectively eliminate immunoreactive T cells)

RN 174230-05-8 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl-

REFERENCE COUNT: 78 THERE ARE 78 CITED REFERENCES AVAILABLE FOR THIS

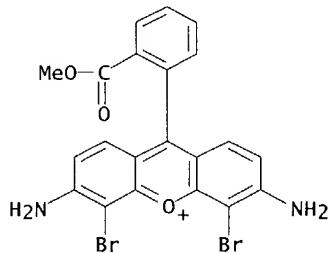
## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 5 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2002:335895 HCPLUS  
 DOCUMENT NUMBER: 137:75306  
 TITLE: Prevention of graft-versus-host disease while preserving graft-versus-leukemia effect after selective depletion of host-reactive T cells by photodynamic cell purging process  
 AUTHOR(S): Chen, Benny J.; Cui, Xiuyu; Liu, Congxiao; Chao, Nelson J.  
 CORPORATE SOURCE: Bone Marrow Transplantation Program, Duke University Medical Center, Durham, NC, 27705, USA  
 SOURCE: Blood (2002), 99(9), 3083-3088  
 CODEN: BLO0AW; ISSN: 0006-4971  
 PUBLISHER: American Society of Hematology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB In this study, we investigated the possibility of selective depletion of donor alloantigen-specific T cells from C57BL/6 (H-2b) mice to prevent graft-vs.-host disease (GVHD). These cells were first activated with irradiated BALB/c (H-2d) host spleen cells in a 5-day mixed lymphocyte culture. Following this activation, a photoactive rhodamine deriv. called 4,5-dibromorhodamine 123 (TH9402), was added. This compd. is selectively retained in the mitochondria of activated host-reactive cells but not tumor-or third-party-specific resting cells. The treated cells were subsequently exposed to visible light (514 nm) to deplete the TH9402-enriched activated host-reactive cells. Treatment with photodynamic cell purging process (PDP) inhibited antihost responses measured by cytotoxic T lymphocytes (CTL) by 93%, and interferon-.gamma. prodn. by 66%. By contrast, anti-BCL1 (BALB/c-origin leukemia/lymphoma) and anti-third-party C3H/HeJ (H-2k) responses were preserved. PDP-treated primed C57BL/6 cells were further tested in vivo. All lethally irradiated BALB/c mice inoculated with BCL1 cells and T-cell-depleted bone marrow cells developed leukemia by day +30, with 50% mortality by 100 days. All mice died of GVHD after addn. of 5 .times. 10<sup>6</sup> untreated primed C57BL/6 cells. However, addn. of same nos. of PDP-treated cells allowed 90% of the recipients to survive more than 100 days without detectable BCL1 tumor cells and free of GVHD. Moreover, PDP-treated primed C57BL/6 cells retained the ability to induce GVHD in the third-party C3H/HeJ mice. These data suggest that PDP can selectively deplete host alloantigen-specific T cells for GVHD prevention and immune and antileukemia function preserve.

IT 174230-05-8, TH9402  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (prevention of graft-vs.-host disease while preserving  
 graft-vs.-leukemia effect after selective depletion of host-reactive T  
 cells by photodynamic cell purging)

RN 174230-05-8 HCPLUS  
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 6 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2001:265273 HCAPLUS  
 DOCUMENT NUMBER: 134:292146  
 TITLE: Rhodamine derivatives for photodynamic diagnosis and treatment  
 INVENTOR(S): Roy, Denis-Claude; Guimond, Martin; Molfino, Nestor A.  
 PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital Maisonneuve-Rosemont  
 SOURCE: PCT Int. Appl., 60 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

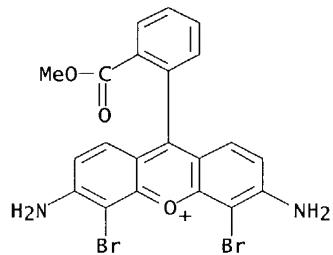
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
BR 2000014135	A	20020521	BR 2000-14135	20001003
EP 1267931	A1	20030102	EP 2000-965683	20001003
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL				
JP 2003510372	T2	20030318	JP 2001-527823	20001003
PRIORITY APPLN. INFO.:			US 1999-157790P	P 19991005
			WO 2000-CA1142	W 20001003

AB The present invention relates to the use of the photoactivatable derivs. for the photodynamic treatment for the selective destruction and/or inactivation of immunol. reactive cells without affecting the normal cells and without causing systemic toxicity for the patient, wherein appropriate intracellular levels of said derivs. are achieved and irradn. of a suitable wavelength and intensity is applied. Examples are given of the selective phototoxicity of rhodamine derivs. against K562 cells, CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and human B cells. Immunol. disorders, including graft-vs-host disease are treated with photodynamic therapy.

IT 333957-97-4  
 RL: BAC (Biological activity or effector, except adverse); BPR (Biological

process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (rhodamine derivs. for photodynamic diagnosis and treatment of immunol.  
 disorders)

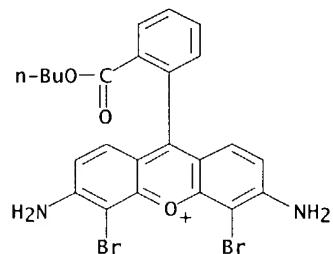
RN 333957-97-4 HCPLUS  
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

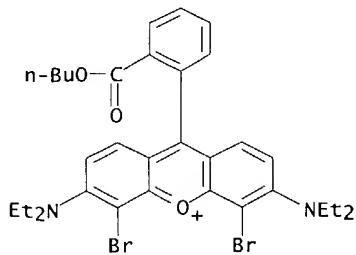
IT 333957-95-2 333957-96-3  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (rhodamine derivs. for photodynamic diagnosis and treatment of immunol.  
 disorders)

RN 333957-95-2 HCPLUS  
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br<sup>-</sup>

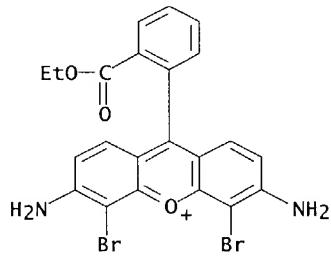
RN 333957-96-3 HCPLUS  
 CN Xanthylium, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-, chloride (9CI) (CA INDEX NAME)

● Cl<sup>-</sup>

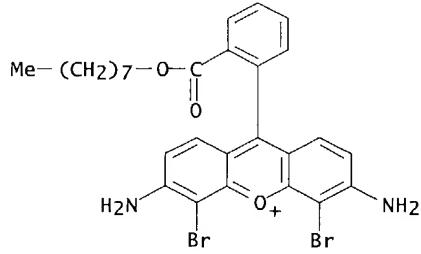
IT 333957-98-5 333957-99-6

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (rhodamine derivs. for photodynamic diagnosis and treatment of immunol.  
 disorders)

RN 333957-98-5 HCPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(ethoxycarbonyl)phenyl]-, bromide  
 (9CI) (CA INDEX NAME)● Br<sup>-</sup>

RN 333957-99-6 HCPLUS

CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-[(octyloxy)carbonyl]phenyl]-,  
 bromide (9CI) (CA INDEX NAME)● Br<sup>-</sup>

REFERENCE COUNT:

8

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS

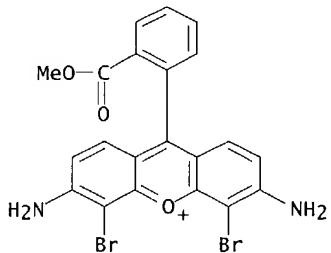
## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 7 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2001:20398 HCPLUS  
 DOCUMENT NUMBER: 134:204394  
 TITLE: Eradication of multiple myeloma and breast cancer  
 cells by TH9402-mediated photodynamic therapy:  
 implication for clinical ex vivo purging of autologous  
 stem cell transplants  
 AUTHOR(S): Brasseur, N.; Menard, I.; Forget, A.; El Jastimi, R.;  
 Hamel, R.; Molfino, N. A.; Van Lier, J. E.  
 CORPORATE SOURCE: Department of Nuclear Medicine and Radiobiology,  
 Faculty of Medicine, Universite de Sherbrooke,  
 Sherbrooke, QC, J1H 5N4, Can.  
 SOURCE: Photochemistry and Photobiology (2000), 72(6), 780-787  
 CODEN: PHCBAP; ISSN: 0031-8655  
 PUBLISHER: American Society for Photobiology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB High-dose chemotherapy combined with autologous transplantation using bone marrow or peripheral blood-derived stem cells (PBSC) is now widely used in the treatment of hematol. malignancies as well as some solid tumors like breast cancer (BC). However, some controversial results were recently obtained in the latter case. The presence of malignant cells in the autograft has been assocd. with the recurrence of the disease, and purging procedures are needed to eliminate this risk. The aim of this study was to evaluate the potential of the photosensitizer 4,5-dibromorhodamine Me ester (TH9402), a dibrominated rhodamine deriv., to eradicate multiple myeloma (MM) and BC cell lines, while sparing more than 50% of normal pluripotential blood stem cells from healthy volunteers. The human BC MCF-7 and T-47D and MM RPMI 8226 and NCI-H929 cell lines were used to optimize the photodynamic purging process. Cell concn. and the cell suspension thickness as well as the dye and light doses were varied in order to eventually treat 1-2 L of apheresis. The light source consisted of two fluorescent scanning tubes emitting green light centered about 515 nm. The cellular uptake of TH9402 was measured during the incubation and washout periods and after photodynamic treatment (PDT) using spectrofluorometric anal. The limiting diln. assay showed that an eradication rate of more than 5 logs is obtained when using a 40 min incubation with 5-10 .mu.M dye followed by a 90 min washout period and a light dose of 5-10 J/cm<sup>2</sup> (2.8 mW/cm<sup>2</sup>) in all cell lines. Agitating the 2 cm thick cell suspension contg. 20 .times. 10<sup>6</sup> cells/mL during PDT was essential for maximal photoinactivation. Expts. on mobilized PBSC obtained from healthy volunteers showed that even more drastic purging conditions than those found optimal for maximal eradication of the malignant cell lines were compatible with a good recovery of hematopoietic progenitors cells. The absence of significant toxicity towards normal hematopoietic stem cells, combined with the 5 logs eradication of cancer cell lines induced by this procedure suggests that TH9402 offers an excellent potential as an ex vivo photodynamic purging agent for autologous transplantation in MM and BC treatment.

IT 174230-05-8, TH9402  
 RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  
 (myeloma and breast cancer cells eradication by TH9402-mediated photodynamic therapy: implication for clin. ex vivo purging of autologous stem cell transplants)

RN 174230-05-8 HCPLUS  
 CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

REFERENCE COUNT: 59 THERE ARE 59 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 8 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 2000:911534 HCPLUS

DOCUMENT NUMBER: 134:66121

TITLE: Compositions and methods for assaying subcellular conditions and processes using energy transfer for drug screening

INVENTOR(S): Dykens, James A.; Velicelebi, Gonul; Ghosh, Soumitra S.

PATENT ASSIGNEE(S): Mitokor, USA

SOURCE: PCT Int. Appl., 189 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

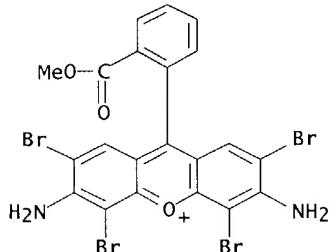
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000079274	A2	20001228	WO 2000-US17380	20000622
WO 2000079274	A3	20020110		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
US 6323039	B1	20011127	US 1999-338122	19990622
US 6280981	B1	20010828	US 2000-514569	20000223
EP 1210596	A2	20020605	EP 2000-943119	20000622
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
JP 2003506014	T2	20030218	JP 2001-505191	20000622
PRIORITY APPLN. INFO.:			US 1999-140433P	P 19990622
			US 1999-338122	A 19990622
			US 2000-176383P	P 20000114
			WO 2000-US17380	W 20000622

AB The invention provides compns. and methods for monitoring subcellular compartments such as organelles by energy transfer techniques that do not require specific intermol. affinity binding events between energy transfer donor and energy transfer acceptor mols. pH. Provided are methods for assaying cellular membrane potential, including mitochondrial membrane potential, by energy transfer methodologies including fluorescence resonance energy transfer (FRET). Diagnostic and drug screening assays

IT are also provided.  
 IT 83796-96-7, Tetrabromo-rhodamine 123  
 RL: ARG (Analytical reagent use); BSU (Biological study, unclassified);  
 ANST (Analytical study); BIOL (Biological study); USES (Uses)  
 (tetrabromorhodamine 123; compns. and methods for assaying subcellular  
 conditions and processes using energy transfer for drug screening)  
 RN 83796-96-7 HCPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,  
 chloride (9CI) (CA INDEX NAME)



● C1 -

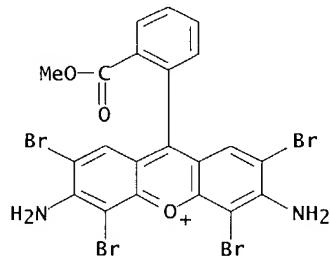
L29 ANSWER 9 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2000:507144 HCPLUS  
 DOCUMENT NUMBER: 133:360498  
 TITLE: Nonthermal ureteral tissue bonding: comparison of  
 photochemical collagen crosslinking with thermal laser  
 bonding  
 AUTHOR(S): Merguerian, Paul A. M. D.; Pugach, Jeff L. M. D.;  
 Lilge, Lothar D.  
 CORPORATE SOURCE: Urology Div., Hospital for Sick Children, Univ. of  
 Toronto, Toronto, ON, Can.  
 SOURCE: Proceedings of SPIE-The International Society for  
 Optical Engineering (1999), 3590(Lasers in Surgery:  
 Advanced Characterization, Therapeutics, and Systems  
 IX), 194-202  
 CODEN: PSISDG; ISSN: 0277-786X  
 PUBLISHER: SPIE-The International Society for Optical Engineering  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB Because of difficulties with suture placement during minimally invasive procedures, many have sought alternative methods of creating tissue anastomoses. Although well studied, thermal laser tissue bonding has the potential of causing collateral thermal injury. Non-thermal tissue bonding agents, which cross-link proteins when activated with light, are currently being explored. We recently reported successful non-thermal bonding using tetrabromorhodamine (TBR). The bond was stronger than sutured repairs but weaker than laser thermal bonding. We currently report our ex-vivo experience with an alternate agent, riboflavin-5-phosphate and compare these results to thermal bonding and TBR. Successful ex vivo photochem. tissue welding with riboflavin of the rabbit ureter was achieved, without the generation of heat. Bond strength similar to that obtained with thermal welding was achieved.

IT 83796-96-7, Tetrabromo-rhodamine 123  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (nonthermal ureteral tissue bonding: comparison of photochem. collagen crosslinking with thermal laser bonding)

RN 83796-96-7 HCPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

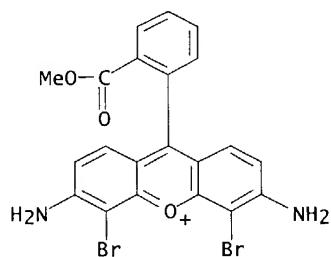


● C1-

REFERENCE COUNT: 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 10 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1999:570227 HCPLUS  
 DOCUMENT NUMBER: 131:308414  
 TITLE: Ex vivo photodynamic purging in chronic myelogenous leukemia and other neoplasias with rhodamine derivatives  
 AUTHOR(S): Villeneuve, Luc  
 CORPORATE SOURCE: Theratechnologies Inc., Montreal, QC, H3B 1S6, Can.  
 SOURCE: Biotechnology and Applied Biochemistry (1999), 30(1), 1-17  
 CODEN: BABIEC; ISSN: 0885-4513  
 PUBLISHER: Portland Press Ltd.  
 DOCUMENT TYPE: Journal; General Review  
 LANGUAGE: English  
 AB A review with 294 refs. Photodynamic therapy (PDT), a cancer treatment already used early in this century, has distinctive advantages over conventional chemotherapy, namely its often obsd. preferential accumulation in cancer cells and its low intrinsic toxicity. Aggressive therapeutic modalities using high doses of chemotherapy and/or radiation therapy are now commonplace treatments for leukemia, lymphoma and various non-haematol. malignancies. These intensive approaches have often been used in assocn. with hematopoietic-progenitor-cell support and have induced major responses and remissions in patients with relapsed and refractory diseases, ultimately contributing to improve the disease-free survival of patients with high risk. This has encouraged Theratechnologies, a Montreal-based pharmaceutical company, to develop photodynamic ex vivo purging procedures, including the development of new photosensitizers and irradn. devices for the safe eradication of neoplastic cells from autologous grafts. Our first specific objective, therefore, was to design, synthesize, purify and test photoactive rhodamine derivs. 4,5-Dibromorhodamine 123 (TH9402), a gas and phosphorus coating characteristic of an efficient scanning fluorescent source for extra-corporeal PDT using rhodamine derivs., was selected because of its photophys. properties, low toxicity and stability. TH9402 photodynamic-cell-therapy process conditions recognized as safe for normal human hematopoietic stem cells and progenitors demonstrated the efficacy of the purging procedure on various leukemias (including chronic-myelogenous-leukemia) as well as non-Hodgkin-leukemias and metastatic-breast-cancer cell lines.  
 IT 174230-05-8, TH 9402  
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (TH 9402; ex vivo photodynamic purging in chronic myelogenous leukemia and other neoplasias with rhodamine derivs.)

RN 174230-05-8 HCPLUS  
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl<sup>-</sup>

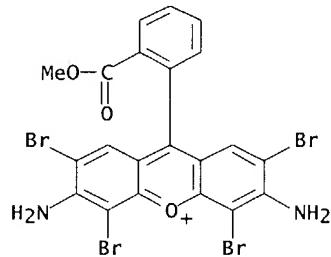
REFERENCE COUNT: 292 THERE ARE 292 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L29 ANSWER 11 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1999:244672 HCPLUS  
 DOCUMENT NUMBER: 130:277634  
 TITLE: Screening for oligonucleotide inhibitors of gene expression using conjugates with activatable reactive substances  
 INVENTOR(S): Prescott, Catherine Denise  
 PATENT ASSIGNEE(S): Smithkline Beecham Corporation, USA  
 SOURCE: PCT Int. Appl., 21 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918116	A1	19990415	WO 1998-US21052	19981007
W: CA, JP, US			W: CA, JP, US	
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE			RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE	
EP 1023312	A1	20000802	EP 1998-952108	19981007
R: BE, CH, DE, DK, FR, GB, IT, LI, NL			R: BE, CH, DE, DK, FR, GB, IT, LI, NL	
JP 2001519141	T2	20011023	JP 2000-514925	19981007
US 6387703	B1	20020514	US 2000-529095	20000406
PRIORITY APPLN. INFO.:			US 1997-61218P	P 19971007
			WO 1998-US21052	W 19981007

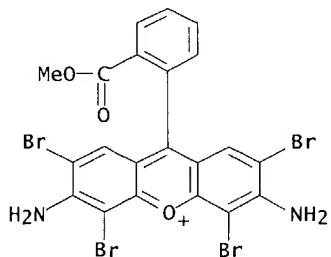
AB A method of screening for compds., particularly oligonucleotides, that modulate gene expression, particularly those which lower gene expression is described. The method uses a conjugate of the oligonucleotide and an activatable reactive group, such as a photosensitizing dye, preferably a compd. that generates reactive oxygen species. Target cells are incubated with the conjugate and the reactive group is activated and the effect on gene expression is assayed.  
 IT 83796-96-7D, Tetrabromo-rhodamine 123, derivs., conjugates with oligonucleotides  
 RL: BUU (Biological use, unclassified); RCT (Reactant); BIOL (Biological study); RACT (Reactant or reagent); USES (Uses)  
 (tetrabromorhodamine 123, photoactivatable inhibition of gene expression using; screening for oligonucleotide inhibitors of gene expression using conjugates with activatable reactive substances)

RN 83796-96-7 HCAPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,  
 chloride (9CI) (CA INDEX NAME)

● Cl<sup>-</sup>

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

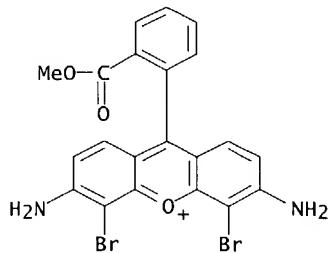
L29 ANSWER 12 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1998:298888 HCAPLUS  
 DOCUMENT NUMBER: 129:51463  
 TITLE: [125I/127I/131I]Iodorhodamine: Synthesis, Cellular Localization, and Biodistribution in Athymic Mice Bearing Human Tumor Xenografts and Comparison with [99mTc]Hexakis(2-methoxyisobutylisonitrile)  
 AUTHOR(S): Harapanhalli, Ravi S.; Roy, Aloka M.; Adelstein, S. James; Kassis, Amin I.  
 CORPORATE SOURCE: Department of Radiology (Nuclear Medicine), Harvard Medical School, Boston, MA, 02115, USA  
 SOURCE: Journal of Medicinal Chemistry (1998), 41(12), 2111-2117  
 CODEN: JMCMAR; ISSN: 0022-2623  
 PUBLISHER: American Chemical Society  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB The synthesis of halogenated rhodamine (Rh) derivs. was carried out by controlling the stoichiometry of the halogenating agents, bromine and iodine monochloride. In the no-carrier-added synthesis of radioiodinated rhodamine 123, direct labeling of rhodamine 123 (Rh 123) with Na125I/Na131I required the presence of the oxidant peracetic acid. 125I/131I-Rh 123 was synthesized in modest yields (40-45%). HPLC purifn. sepd. Rh 123 from its mono- and diiodo derivs. Monohalogenation of Rh 123 did not alter the compd.'s ability to permeate viable cells and localize in mitochondria. 125I/131I-Rh 123 was stable in serum in vitro but rapidly metabolized after i.v. injection into mice. Consequently, scintigraphy and biodistribution data reveal poor targeting of s.c. growing human tumor xenografts. The results are compared to those obtained following the administration of [99mTc]hexakis(2-methoxyisobutylisonitrile) which also did not image human tumor xenografts in nude mice.  
 IT 83796-96-7P  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (prep. and biodistribution of radioiodinated rhodamine 123 in tumor imaging)  
 RN 83796-96-7 HCAPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

REFERENCE COUNT: 23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 13 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1997:104125 HCPLUS  
 DOCUMENT NUMBER: 126:183387  
 TITLE: Spectroscopic and photophysical investigations on the nature of localization of rhodamine-123 and its dibromo derivative in different cell lines  
 AUTHOR(S): Villeneuve, Luc; Pal, Prabir; Durocher, Gilles; Migneault, David; Girard, Denis; Giasson, Richard; Balassy, Antonia; Blanchard, Louise; Gaboury, Louis  
 CORPORATE SOURCE: Laboratoire de pathologie moléculaire, Département de pathologie, Université de Montréal, Montréal, QC, H3C 3J7, Can.  
 SOURCE: Journal of Fluorescence (1996), 6(4), 209-219  
 CODEN: JOFLEN; ISSN: 1053-0509  
 PUBLISHER: Plenum  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB Steady-state and time-resolved spectroscopic properties of rhodamine-123 (rh123) and 4,5-dibromorhodamine Me ester (dbr123) bound to different cell lines are evaluated. Studies are also performed on the dye bound to extd. mitochondria. Results are compared with those obtained in homogeneous and microheterogeneous media. Results suggest that these dyes can specifically bind only with cell mitochondria. As a result of binding, excitation and emission spectra are red shifted by 10 to 12 nm. The fluorescence decay of these dyes bound to mitochondria shows two lifetimes. Values are about 4.0 and 2.0 ns for rh123 and about 1.9 and 0.5 ns for dbr123. Detailed global anal. of emission wavelength and dye concn. dependences of the fluorescence decay is performed. Results indicate that these dyes are bound to two different binding sites at mitochondria. The decay-assocd. fluorescence spectrum for the species corresponding to each binding site is recovered. Species 1, corresponding to the longer lifetime, is found to be more red shifted compared to species 2. The fluorescence of species 2 is heavily quenched. The origin of this quenching is explained in terms of resonance energy transfer between donor species 2 and acceptor species 1. The possible nature of the two binding sites is also discussed.  
 IT 174230-05-8  
 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (spectroscopic and photophys. investigations on the nature of localization of rhodamine-123 and its dibromo deriv. in different cell lines)  
 RN 174230-05-8 HCPLUS  
 CN Xanthylum, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

L29 ANSWER 14 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1996:538169 HCPLUS

DOCUMENT NUMBER: 125:260967

TITLE: Spectroscopic and photophysical properties of some new rhodamine derivatives in cationic, anionic and neutral micelles

AUTHOR(S): Pal, P.; Zeng, H.; Durocher, G.; Girard, D.; Giasson, R.; Blanchard, L.; Gaboury, L.; Villeneuve, L.

CORPORATE SOURCE: Laboratoire de photophysique moléculaire, Département de chimie, Université de Montréal, C.P. 6128, Succ. Centre-ville, Montréal, Que., H3C 3J7, Can.

SOURCE: Journal of Photochemistry and Photobiology, A: Chemistry (1996), 98(1-2), 65-72  
CODEN: JPPCEJ; ISSN: 1010-6030

PUBLISHER: Elsevier

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The spectroscopic and photophys. characterization of rhodamine 123 (dye 1), 4,5-dibromorhodamine Me ester (dye 2) and 4,5-dibromorhodamine Bu ester (dye 3) are reported in homogeneous media like water and some alcs. and also in microheterogeneous media; anionic sodium dodecylsulfate (SDS), cationic cetyltrimethylammonium bromide (CTAB) and neutral triton X-100 (TX) micelles. The selective biodistribution of these ionic drugs in tissues and membranes strongly influence their photosensitizing properties which have been part of our earlier studies. Results suggest that the hydrogen bonding capability of the amino end group lone pair of these dyes dominates in water. All these dyes interact with anionic SDS micelles. The interaction is mainly electrostatic in nature. At low SDS concns. (below c.m.c.), dye-SDS aggregate formation takes place. But above c.m.c. only the monomeric dye form is obsd. The penetration of dye 3 in SDS is a little less compared to dyes 1 and 2. Dyes 2 and 3 show a finite interaction with CTAB micelle unlike dye 1. With neutral TX micelles all the dyes form strong complexes. The fluorescence quantum yield (.PHI.F) of these three dyes in TX is lower. In time-resolved fluorescence expts., two lifetimes are obsd. The effects of the TX concn. on the fluorescence decay are measured. The decay assocd. spectra of dye 2 in TX are obtained by global compartmental anal. The dye-surfactant interaction mechanisms are also discussed.

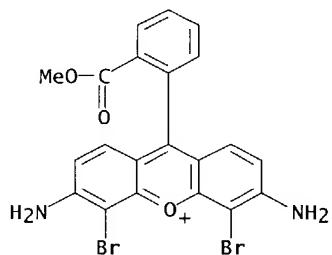
IT 174230-05-8 174230-06-9

RL: PRP (Properties)

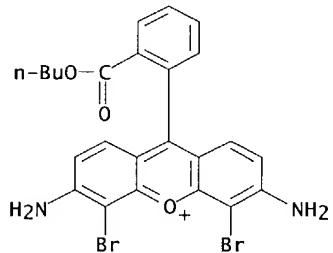
(spectroscopic and photophys. properties of rhodamine derivs. in homogeneous media and micelles)

RN 174230-05-8 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

● Cl<sup>-</sup>

RN 174230-06-9 HCPLUS  
 CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)

● Cl<sup>-</sup>

L29 ANSWER 15 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1996:96151 HCPLUS  
 DOCUMENT NUMBER: 124:197239  
 TITLE: Phototoxicity of some bromine-substituted rhodamine dyes: synthesis, photophysical properties and application as photosensitizers  
 AUTHOR(S): Pal, PRabir; Zeng, Hualing; Durocher, Gilles; Girard, Denis; Li, Tiechao; Gupta, Ajay K.; Giasson, Richard; Blanchard, Louise; Gaboury, Louis; et al.  
 CORPORATE SOURCE: Lab. Photophys. Mol., Univ. Montreal, Montreal, QC, Can.  
 SOURCE: Photochemistry and Photobiology (1996), 63(2), 161-8  
 CODEN: PHCBAP; ISSN: 0031-8655  
 PUBLISHER: American Society for Photobiology  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB The synthesis of some bromine-substituted rhodamine derivs., viz., 4,5-dibromorhodamine Me ester (dye 2) and 4,5-dibromorhodamine Bu ester (dye 3), are reported. These dyes were synthesized to promote a more efficient cancer cell photosensitizer for potential use in in vitro bone marrow purging in prepn. for autologous bone marrow transplantation. Spectroscopic and photophys. characterization of these dyes together with rhodamine 123 (dye 1) are reported in water, methanol, ethanol and also in a microheterogeneous system, sodium dodecyl sulfate. The possible mechanism of photosensitization os characterized in terms of singlet oxygen efficiency of these dyes. Singlet oxygen quantum yields for

bromine-substituted dyes are in the range of 0.3-0.5 depending on the solvent. For dye 1 no singlet oxygen prodn. is found. The photodynamic actions of these dyes in different cell lines are tested. It was found that dye 2 and dye 3 are efficient photosensitizers and mediate eradication of K562, EM2, myeloid cell lines (CML) and the SMF-AI rhabdomyosarcoma line.

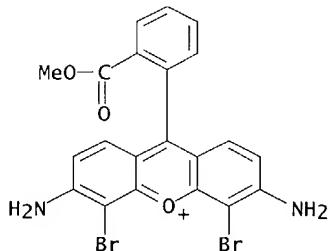
IT 174230-05-8P 174230-06-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(phototoxicity of some bromine-substituted rhodamine dyes: synthesis, photophys. properties and application in leukemia photosensitzations with laser radiation)

RN 174230-05-8 HCPLUS

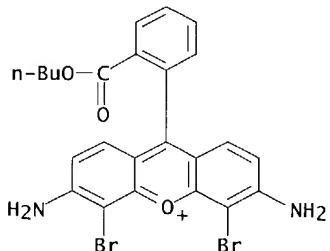
CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1-

RN 174230-06-9 HCPLUS

CN Xanthylium, 3,6-diamino-4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1-

L29 ANSWER 16 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1995:709283 HCPLUS

DOCUMENT NUMBER: 123:164113

TITLE: DMSO affects the efficiency of photolabeling of tetrabrominated rhodamine to collagen fibers.

AUTHOR(S): Jacques, Steven L.; Awazu, Kunio; Hasan, Tayyaba  
CORPORATE SOURCE: M. D. Anderson Cancer Center, Univ. Texas, Houston,  
TX, 77030, USA

SOURCE: Proceedings of SPIE-The International Society for Optical Engineering (1995), 2391(Laser-Tissue Interaction VI), 232-7  
 CODEN: PSISDG; ISSN: 0277-786X

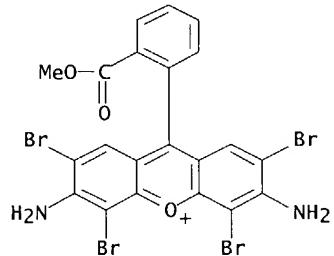
PUBLISHER: SPIE-The International Society for Optical Engineering  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB The ability to photolabel a compd., tetrabrominated rhodamine (TBR), onto collagen gels was tested. The influence of DMSO on the efficiency of photolabeling was detd. DMSO enhances the photolabeling presumably by allowing TBR to become more closely assocd. to the collagen fibers such that upon photon absorption which causes debromination to yield a radical, the radical can covalently link to the collagen.

IT 83796-96-7  
 RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)  
 (DMSO affects the efficiency of photolabeling of tetrabrominated rhodamine to collagen fibers)

RN 83796-96-7 HCPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

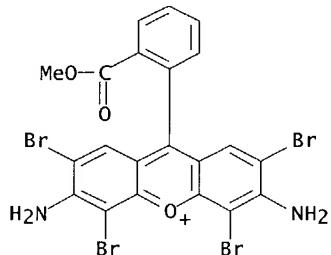
L29 ANSWER 17 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1995:583719 HCPLUS  
 DOCUMENT NUMBER: 123:78614  
 TITLE: Photodynamic therapy with cationic photosensitizers  
 AUTHOR(S): Kessel, David; Woodburn, Kathryn; Chang, CK;  
 Henderson, BW  
 CORPORATE SOURCE: Department Pharmacology, Wayne State University School  
 Medicine, Detroit, MI, USA  
 SOURCE: Proceedings of SPIE-The International Society for  
 Optical Engineering (1995), 2371, 334-8  
 CODEN: PSISDG; ISSN: 0277-786X

DOCUMENT TYPE: Journal  
 LANGUAGE: English

AB We characterized sites of photodamage catalyzed by two cationic photosensitizers, tetrabromo-rhodamine 123 (TBR), which is recognized by the multidrug transporter, and a monocationic porphyrin (MCP) which is not. The transporter is an outward transport system assocd. with examples of drug resistance. Irradn. of multidrug-resistant cells treated with TBR resulted in highly-selective photodamage to the transporter site, while MCP catalyzed nonspecific membrane damage to cells regardless of transporter expression.

IT 83796-96-7, Tetrabromo-rhodamine 123  
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
 (photodynamic therapy with cationic photosensitizers)

RN 83796-96-7 HCAPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl<sup>-</sup>

L29 ANSWER 18 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1995:453091 HCAPLUS  
 DOCUMENT NUMBER: 122:285634  
 TITLE: Selective photodynamic inactivation of a multidrug transporter by a cationic photosensitizing agent  
 AUTHOR(S): Kessel, D; Woodburn, K  
 CORPORATE SOURCE: School of Medicine, Wayne State University, Detroit, MI, 48201, USA  
 SOURCE: British Journal of Cancer (1995), 71(2), 306-10  
 CODEN: BJCAAI; ISSN: 0007-0920

DOCUMENT TYPE: Journal

LANGUAGE: English

AB We have characterized sites of photodamage catalyzed by the cationic photosensitizer tetrabromorhodamine 123, using P388 murine leukemia cells and a subline (P388/ADR) which has a multidrug resistance phenotype and hyperexpresses mdr1 mRNA for P-glycoprotein. Fluorescence emission spectra were consistent with sensitizer localization in hydrophobic regions of the P388 cell, and in more aq. loci in P388/ADR. Subsequent irradn. resulted in photodamage to the P388 cells, resulting in loss of viability. In contrast, P388/ADR cells were unaffected except for an irreversible inhibition of P-glycoprotein, leading to enhanced accumulation of daunorubicin and rhodamine 123 and a corresponding increase in daunorubicin cytotoxicity. These results are consistent with the premise that substrates for P-glycoprotein are confined to membrane loci assocd. with the transporter, and indicate a very limited migration of cytotoxic photoproducts in a cellular environment.

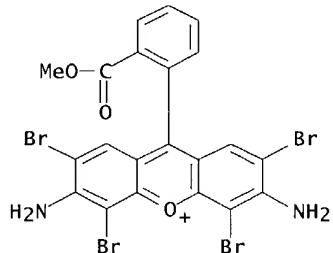
IT 83796-96-7, Tetrabromo-rhodamine 123

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(photodynamic inactivation of multidrug transporter in leukemia cells by cationic photosensitizer tetrabromorhodamine 123 with visible light)

RN 83796-96-7 HCAPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

L29 ANSWER 19 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1993:18589 HCPLUS

DOCUMENT NUMBER: 118:18589

TITLE: Mapping radiant energy distributions during laser  
irradiation of collagen phantoms by photolabeling with  
tetrabrominated rhodamine

AUTHOR(S): Jacques, Steven L.; Hasan, Tayyaba

CORPORATE SOURCE: Laser Biol. Res. Lab., Univ. Texas, Houston, TX,  
77030, USASOURCE: Proceedings of SPIE-The International Society for  
Optical Engineering (1992), 1646(Proc. Laser-Tissue  
Interact. III, 1992), 219-26

CODEN: PSISDG; ISSN: 0277-786X

DOCUMENT TYPE: Journal

LANGUAGE: English

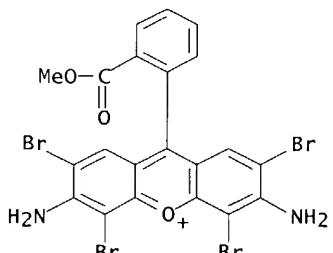
AB A method is proposed for mapping laser light distributions in gel phantoms  
by measurement of the distribution of a fluorescent compd. that has been  
photolabeled to the gel by the laser irradiance. A preliminary study of  
photolabeling by an argon laser using tetrabrominated rhodamine (TBR) was  
conducted in collagen gel phantoms to illustrate the feasibility of the  
method. A discussion of the basic quant. relationships for anal. of  
measurements is presented.

IT 83796-96-7

RL: BIOL (Biological study)

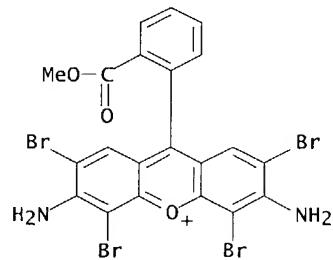
(photolabeling with, in laser radiation energy distribution mapping in  
collagen gel phantoms)

RN 83796-96-7 HCPLUS

CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,  
chloride (9CI) (CA INDEX NAME)

● C1 -

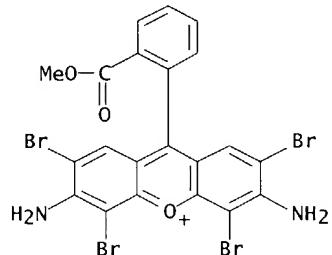
L29 ANSWER 20 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1992:146920 HCPLUS  
 DOCUMENT NUMBER: 116:146920  
 TITLE: A test of the singlet oxygen mechanism of cationic dye photosensitization of mitochondrial damage  
 AUTHOR(S): Bunting, James R.  
 CORPORATE SOURCE: Baylor Res. Inst., Dallas, TX, 75226, USA  
 SOURCE: Photochemistry and Photobiology (1992), 55(1), 81-7  
 CODEN: PHCBAP; ISSN: 0031-8655  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB Arom. cationic dyes have a potential as photo-chemotherapeutic agents because they are selectively concd. into the mitochondria of cancerous cells. The mechanism of cytophototoxicity has been proposed to be primarily due to dye sensitized photogeneration of highly toxic singlet oxygen (102) at the mitochondria. This hypothesis was tested by measuring the relative phototoxicity of a collection of arom. cationic dyes towards respiring rat-liver mitochondria (RLM), upon addn. of 514 nm laser light. The effectiveness of dye photosensitization towards destruction of RLM function was assayed by its effect on the RLM membrane potential. Three phys. parameters of dye phototoxicity were independently measured and a relative phototoxicity calcd. assuming adherence of mechanism in the 102 hypothesis. Quantum yields of dye-sensitized 102 prodn. were estd., either from time-resolved luminescence measurements of photosensitized 102 formed, or by comparing rates of photobleaching of 102 trap; the relative partition of dye into mitochondrial lipid was detd. gravimetrically; and the optical d. of dye was detd. in a lipid like Triton X 100 micellar environment. Under the assumption of the 102 hypothesis, these parameters were used to predict a relative phototoxicity which was compared with that obsd. For 12 of the 14 dyes investigated, the obsd. and predicted phototoxicities were linearly correlated ( $r = 0.85$ ), suggesting support of the 10- and 1000-fold more potent than predicted, suggesting an addnl. factor at play in their phototoxicity.  
 IT 83796-96-7, Tetrabromo-rhodamine 123  
 RL: BIOL (Biological study)  
 (photosensitization by, of liver mitochondria, singlet oxygen mechanism of evaluation in)  
 RN 83796-96-7 HCPLUS  
 CN Xanthylum, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

L29 ANSWER 21 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1989:473949 HCPLUS  
 DOCUMENT NUMBER: 111:73949  
 TITLE: Rhodamine dyes as potential agents for photochemotherapy of cancer in human bladder carcinoma cells

AUTHOR(S): Shea, Christopher R.; Chen, Norah; Wimberly, Joanne;  
 Hasan, Tayyaba  
 CORPORATE SOURCE: Dep. Dermatol., Harvard Med. Sch., Boston, MA, 02114,  
 USA  
 SOURCE: Cancer Research (1989), 49(14), 3961-5  
 CODEN: CNREA8; ISSN: 0008-5472  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB The phototoxicity *in vitro* of rhodamine 123 and tetrabromorhodamine 123 (TBR) was compared in order to assess the photochemotherapeutic potential of these compds. Exposure to 514.5-nm radiation from an Ar ion laser caused phototoxicity in MGH-U1 bladder carcinoma cells previously treated with either dye at 10  $\mu\text{M}$  for 30 min. As assessed by colony formation and cellular morphol., TBR was markedly more phototoxic than rhodamine 123, reflecting increased intersystem crossing of TBR to the triplet manifold via spin-orbital coupling induced by the heavy Br atoms. Photoreactions of TBR very efficiently generated singlet O (102) in soln.; furthermore, irradn. of TBR-treated cells was significantly more toxic when performed in the presence of deuterium oxide, an enhancer of damage caused by 102. Retention of fluorescence in TBR-treated cells was enhanced by irradn., indicating that a stable photoproduct may be formed in reaction with cellular components.  
 IT 83796-96-7  
 RL: PRP (Properties)  
 (phototoxicity of, photochemotherapy of human bladder carcinoma in relation to)  
 RN 83796-96-7 HCPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

L29 ANSWER 22 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1989:208583 HCPLUS  
 DOCUMENT NUMBER: 110:208583  
 TITLE: Phototoxicity of rhodamine dyes  
 AUTHOR(S): Shea, Christopher R.; Chen, Norah; Hasan, Tayyaba  
 CORPORATE SOURCE: Massachusetts Gen. Hosp., Harvard Med. Sch., Boston,  
 MA, 02114, USA  
 SOURCE: Proceedings of SPIE-The International Society for  
 Optical Engineering (1989), 997(Adv. Photochemother.),  
 48-57  
 CODEN: PSISDG; ISSN: 0277-786X  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 AB Rhodamine-123 (R123) and tetrabromo-R123 (TBR) have been evaluated *in vitro* as potential photosensitizers with laser radiation. R123 localizes selectively in mitochondria of MGH-U1 bladder carcinoma cells exposed to 10  $\mu\text{M}$  R123 for 30 min, and under these conditions R123 is a weak photosensitizer. Incubation with R123 for longer times enhances its

phototoxicity, and is assocd. with a modification of its intracellular localization. TBR is .apprx.100-fold more phototoxic than R123, as assessed either by [<sup>3</sup>H]thymidine uptake or vital staining. Actively proliferating cells are more sensitive to either R123 or TBR phototoxicity than are plateau-phase, confluent cultures.

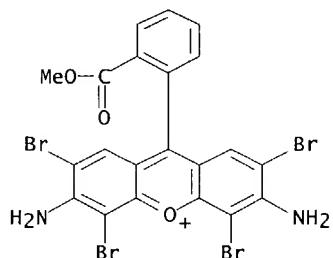
IT 83796-96-7

RL: PRP (Properties)

(phototoxicity of, to bladder carcinoma cells with laser radiation)

RN 83796-96-7 HCPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● C1 -

L29 ANSWER 23 OF 24 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1983:211699 HCPLUS

DOCUMENT NUMBER: 98:211699

TITLE: Triplet anisotropy decay measurements of DNA internal motion

AUTHOR(S): Hogan, Michael; Wang, Johnny; Austin, R. H.

CORPORATE SOURCE: Dep. Biochem. Sci., Princeton Univ., Princeton, NJ, 08540, USA

SOURCE: Ciba Foundation Symposium (1983), 93(Mobility Funct. Proteins Nucleic Acids), 226-45

CODEN: CIBSB4; ISSN: 0300-5208

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Triplet anisotropy decay techniques were used to measure the internal flexibility and overall rotational motions of DNA over a time range of 15 ns to 200 .mu.s. Nearly monodisperse DNA fragments with lengths varying from 65-600 base pairs were studied with the intercalating dye methylene blue as a triplet probe. The slow end-over-end tumbling of short DNA fragments (<165 base pairs) is as predicted for a rigid rod. A longer DNA fragment (600 base pairs) experiences slow segmental motions of its helix axis. At the earliest times, anisotropy decays more rapidly than expected for a rigid rod, suggesting that, when it is bound, methylene blue monitors fast internal motions of the helix. Since the rodlike end-over-end tumbling rules out fast bending motions (for short DNA fragments), the fast components of DNA anisotropy decay must be due to twisting motions of the helix, occurring with a time const. of .apprx.50 ns. The same techniques were used to measure the conformational flexibility of DNA in the nucleosome. It is concluded that, when the DNA helix is wrapped to form a nucleosome, it experiences substantial internal flexibility, occurring with a time const. of .apprx.30 ns. The amplitude and time-scale of this motion are similar to that seen in the uncomplexed DNA helix.

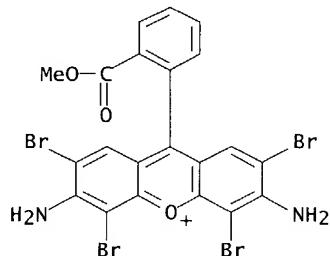
IT 83796-96-7

RL: BIOL (Biological study)

(triplet anisotropy decay of, in DNA, internal motions in relation to)

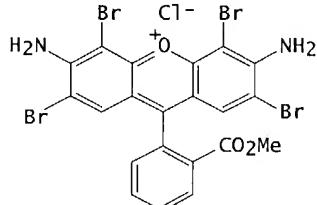
RN 83796-96-7 HCPLUS

CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-, chloride (9CI) (CA INDEX NAME)



● Cl-

L29 ANSWER 24 OF 24 HCAPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 1983:1816 HCAPLUS  
 DOCUMENT NUMBER: 98:1816  
 TITLE: DNA motions in the nucleosome core particle  
 AUTHOR(S): Wang, J.; Hogan, M.; Austin, R. H.  
 CORPORATE SOURCE: Dep. Biochem. Sci., Princeton Univ., Princeton, NJ, 08544, USA  
 SOURCE: Proceedings of the National Academy of Sciences of the United States of America (1982), 79(19), 5896-900  
 CODEN: PNASA6; ISSN: 0027-8424  
 DOCUMENT TYPE: Journal  
 LANGUAGE: English  
 GI



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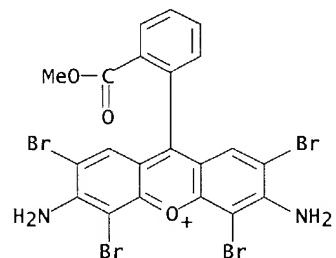
AB Time-resolved triplet-state anisotropy decay techniques employing the intercalating agents, methylene blue and tetrabromorhodamine 123 chloride (I), the latter prep'd. by bromination of rhodamine 123, were used to measure the conformational flexibility of DNA in the chicken erythrocyte nucleosome. In a nucleosome, the DNA helix experiences substantial internal flexibility, which occurs with a time const. of .apprx.30 ns. The data can be fit well by a modified version of the Barkley-Zimm model for DNA motion, allowing only DNA twisting motions and the overall tumbling of the nucleosome. That fit yields a calcd. torsional rigidity equal to 1.8 .times. 10<sup>-19</sup> erg-cm, a value equal to that measured for uncomplexed DNA. Such similarity suggests that large, fast twisting motions of the DNA helix persist, nearly unaltered, when DNA is wrapped to form a nucleosome.

IT 83796-96-7P  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (prep'n. of, as DNA intercalating agent)

RN 83796-96-7 HCAPLUS  
 CN Xanthylium, 3,6-diamino-2,4,5,7-tetrabromo-9-[2-(methoxycarbonyl)phenyl]-,

ROY 10/088,072

chloride (9CI) (CA INDEX NAME)



● Cl<sup>-</sup>

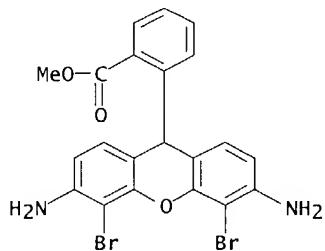
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L39 10 SEA FILE=REGISTRY ABB=ON PLU=ON (13558-31-1/BI OR 177989-33-2  
/BI OR 177989-34-3/BI OR 177989-35-4/BI OR 177989-36-5/BI OR  
177989-37-6/BI OR 177989-38-7/BI OR 62669-70-9/BI OR 71-36-3/BI  
OR 81-88-9/BI)  
L62 2 SEA FILE=REGISTRY ABB=ON PLU=ON "4,5-DIBROMO" AND "BIS(DIETHY  
LAMINO)"  
L63 5 SEA FILE=REGISTRY ABB=ON PLU=ON L39 AND BR=2  
L64 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L62  
L65 2 SEA FILE=HCAPLUS ABB=ON PLU=ON L63  
L66 3 SEA FILE=HCAPLUS ABB=ON PLU=ON (L64 OR L65)

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L66 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN  
ACCESSION NUMBER: 2002:777919 HCAPLUS  
DOCUMENT NUMBER: 137:280622  
TITLE: Halogenated rhodamine dye derivatives and their  
therapeutic applications  
INVENTOR(S): Habi, Abdelkrim; Gravel, Denis; Villeneuve, Luc;  
Forte, Jean-Pierre; Su, Hongsheng; Vaillancourt, Marc  
PATENT ASSIGNEE(S): Theratechnologies Inc., Can.  
SOURCE: PCT Int. Appl., 117 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

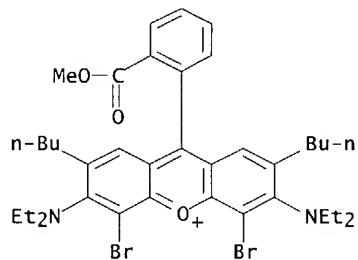
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002079183	A1	20021010	WO 2002-CA438	20020327
WO 2002079183	C1	20030220		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
EP 1276734	A1	20030122	EP 2002-708105	20020327
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
BR 2002004489	A	20030401	BR 2002-4489	20020327
US 2003212126	A1	20031113	US 2003-297088	20030530
CA 2001-2342675 A 20010402				
US 2001-822223 A 20010402				
WO 2002-CA438 W 20020327				

PRIORITY APPLN. INFO.: MARPAT 137:280622  
AB Bromo derivs. of rhodamine 110, rhodamine B, and rhodamine 6G and other  
halo rhodamine derivs. are useful as intermediates and as bactericides and  
antiviral agents and in the treatment of immunol. disorders. In an  
example, rhodamine B Me ester was dihydrogenated and then brominated and  
oxidized and treated with acetic acid to provide the purple acetate salt  
of 2,7-dibromorhodamine B Me ester.  
IT 177989-33-2  
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(bacteriostatic agent; halogenated rhodamine dye derivs. and their  
therapeutic applications)  
RN 177989-33-2 HCAPLUS  
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, methyl ester  
(9CI) (CA INDEX NAME)



IT 467232-23-1P  
 RL: IMF (Industrial manufacture); TEM (Technical or engineered material use); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation);  
 USES (Uses)  
 (red dye; prodn. of halogenated rhodamine dye derivs. and their therapeutic applications)

RN 467232-23-1 HCPLUS  
 CN Xanthylum, 4,5-dibromo-2,7-dibutyl-3,6-bis(diethylamino)-9-[2-(methoxycarbonyl)phenyl]-, bromide (9CI) (CA INDEX NAME)



● Br-

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L66 ANSWER 2 OF 3 HCPLUS COPYRIGHT 2004 ACS on STN  
 ACCESSION NUMBER: 2001:265273 HCPLUS  
 DOCUMENT NUMBER: 134:292146  
 TITLE: Rhodamine derivatives for photodynamic diagnosis and treatment  
 INVENTOR(S): Roy, Denis-Claude; Guimond, Martin; Molfino, Nestor A.  
 PATENT ASSIGNEE(S): Universite de Montreal, Can.; Hopital Maisonneuve-Rosemont  
 SOURCE: PCT Int. Appl., 60 pp.  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2001024824	A1	20010412	WO 2000-CA1142	20001003
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,				

SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,  
 YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM  
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,  
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,  
 CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

BR 2000014135 A 20020521 BR 2000-14135 20001003

EP 1267931 A1 20030102 EP 2000-965683 20001003

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, SI, LT, LV, FI, RO, MK, CY, AL

JP 2003510372 T2 20030318 JP 2001-527823 20001003

PRIORITY APPLN. INFO.: US 1999-157790P P 19991005  
 WO 2000-CA1142 W 20001003

AB The present invention relates to the use of the photoactivable derivs. for the photodynamic treatment for the selective destruction and/or inactivation of immunol. reactive cells without affecting the normal cells and without causing systemic toxicity for the patient, wherein appropriate intracellular levels of said derivs. are achieved and irradn. of a suitable wavelength and intensity is applied. Examples are given of the selective phototoxicity of rhodamine derivs. against K562 cells, CEM cells, PHA-activated lymphocytes, activated CD4+ and CD8+ cells and human B cells. Immunol. disorders, including graft-vs-host disease are treated with photodynamic therapy.

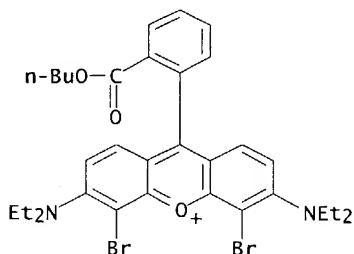
IT 333957-96-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(rhodamine derivs. for photodynamic diagnosis and treatment of immunol. disorders)

RN 333957-96-3 HCPLUS

CN Xanthylum, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-, chloride (9CI) (CA INDEX NAME)



● C1 -

REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L66 ANSWER 3 OF 3 HCPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER: 1996:379805 HCPLUS

DOCUMENT NUMBER: 125:52522

TITLE: Novel rhodamine derivatives for photodynamic therapy of cancer and in vitro purging of the leukemias

INVENTOR(S): Gaboury, Louis; Giasson, Richard; Li, Tiechao; Gupta, Ajay Kumar; Villeneuve, Luc

PATENT ASSIGNEE(S): Universite De Montreal, Can.

SOURCE: PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9607431	A1	19960314	WO 1995-CA485	19950816
W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TT RW: KE, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG				
US 5556992	A	19960917	US 1994-300179	19940902
CA 2197435	AA	19960314	CA 1995-2197435	19950816
AU 9532488	A1	19960327	AU 1995-32488	19950816
AU 688100	B2	19980305		
EP 773794	A1	19970521	EP 1995-928907	19950816
EP 773794	B1	20010620		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE BR 9508779 A 19971223 BR 1995-8779 19950816 JP 10505349 T2 19980526 JP 1995-509057 19950816 AT 202286 E 20010715 AT 1995-928907 19950816 ES 2160173 T3 20011101 ES 1995-928907 19950816 PT 773794 T 20011228 PT 1995-95928907 19950816 US 5773460 A 19980630 US 1996-674247 19960701 GR 3036636 T3 20011231 GR 2001-401494 20010917				
PRIORITY APPLN. INFO.: US 1994-300179 A 19940902 WO 1995-CA485 W 19950816				

AB The present invention relates to novel photoactivable rhodamine derivs. for enhancing high quantum-yield prodn. and singlet oxygen generation upon irradn. with light while maintaining desirable differential retention of rhodamine between normal and cancer cells, said derivs. are selected from the group consisting of 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Me ester hydrochloride); 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Et ester hydrochloride); 4,5-dibromorhodamine 123 (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid octyl ester hydrochloride); 4,5-dibromorhodamine 110 Bu ester (2-(4,5-dibromo-6-amino-3-imino-3H-xanthen-9-yl)-benzoic acid Bu ester hydrochloride); rhodamine B Bu ester (2-(6-Et amino-3-Et imino-3H-xanthen-9-yl)-benzoic acid Bu ester hydrochloride); and photoactivable derivs. thereof; whereby photoactivation of the derivs. induces cell killing while unactivated derivs. are substantially non-toxic to cells. Also, the present invention relates to the use of the photoactivable derivs. of the present invention for the photodynamic therapy of a cancer patient by destroying human cancer cells, wherein appropriate intracellular levels of the derivs. are achieved and irradn. with light of a suitable wavelength is applied. The present invention also relates to a method for the photodynamic therapy of a patient suffering from leukemias, disseminated multiple myelomas or lymphomas.

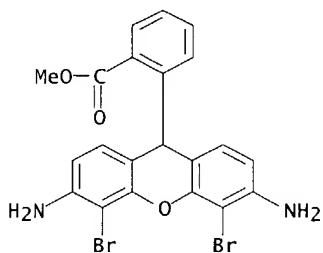
IT 177989-33-2 177989-34-3 177989-35-4

177989-36-5 177989-37-6

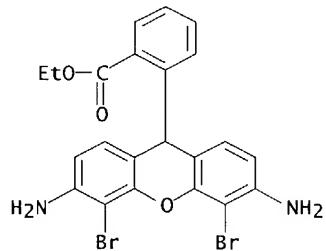
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(rhodamine derivs. for photodynamic therapy of cancer and leukemias)

RN 177989-33-2 HCPLUS

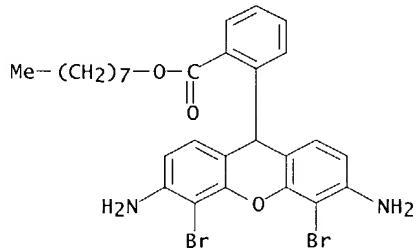
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, methyl ester (9CI) (CA INDEX NAME)



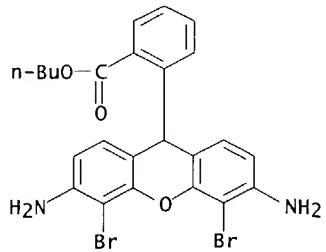
RN 177989-34-3 HCPLUS  
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, ethyl ester  
(9CI) (CA INDEX NAME)



RN 177989-35-4 HCPLUS  
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, octyl ester  
(9CI) (CA INDEX NAME)

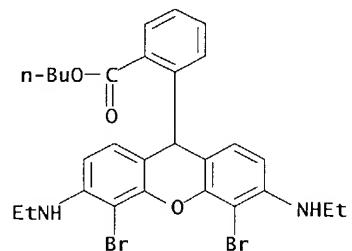


RN 177989-36-5 HCPLUS  
CN Benzoic acid, 2-(3,6-diamino-4,5-dibromo-9H-xanthen-9-yl)-, butyl ester  
(9CI) (CA INDEX NAME)



ROY 10/088,072

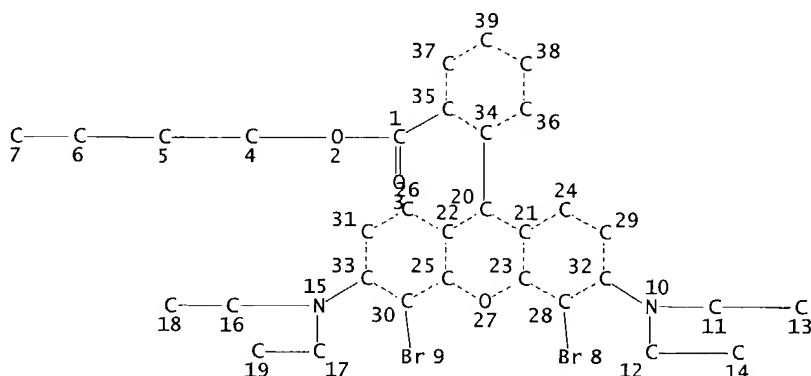
RN 177989-37-6 HCPLUS  
CN Benzoic acid, 2-[4,5-dibromo-3,6-bis(ethylamino)-9H-xanthen-9-yl]-, butyl ester (9CI) (CA INDEX NAME)



ROY 10/088,072

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## NODE ATTRIBUTES:

DEFINITION: DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

#### **GRAPH ATTRIBUTES:**

RING(S) ARE ISOLATED OR EMBEDDED

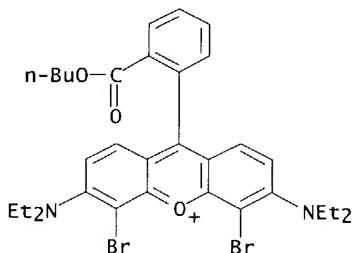
NUMBER OF NODES IS 39

STEREO ATTRIBUTES: NONE

L77 1 SEA FILE=REGISTRY FAM FUL L75

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L77 1 ANSWERS REGISTRY COPYRIGHT 2004 ACS on STN  
IN Xanthylium, 4,5-dibromo-9-[2-(butoxycarbonyl)phenyl]-3,6-bis(diethylamino)-  
, chloride (9CI)  
MF C32 H37 Br2 N2 O3 . Cl



• C1 -

ALL ANSWERS HAVE BEEN SCANNED